

L1           STRUCTURE UPLOADED

=> d l1

L1 HAS NO ANSWERS

L1           STR

\* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT \*

Structure attributes must be viewed using STN Express query preparation.

=> s l1

SAMPLE SEARCH INITIATED 15:08:04 FILE 'REGISTRY'

SAMPLE SCREEN SEARCH COMPLETED -           3 TO ITERATE

100.0% PROCESSED           3 ITERATIONS

3 ANSWERS

SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS:   ONLINE   \*\*COMPLETE\*\*

BATCH   \*\*COMPLETE\*\*

PROJECTED ITERATIONS:           3 TO       163

PROJECTED ANSWERS:           3 TO       163

L2           3 SEA SSS SAM L1

=> s l1 ful

FULL SEARCH INITIATED 15:08:08 FILE 'REGISTRY'

FULL SCREEN SEARCH COMPLETED -           92 TO ITERATE

100.0% PROCESSED           92 ITERATIONS

81 ANSWERS

SEARCH TIME: 00.00.01

L3           81 SEA SSS FUL L1

=> file caplus

COST IN U.S. DOLLARS

SINCE FILE

TOTAL

ENTRY

SESSION

FULL ESTIMATED COST

166.94

167.15

FILE 'CAPLUS' ENTERED AT 15:08:11 ON 28 APR 2006

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FILE COVERS 1907 - 28 Apr 2006 VOL 144 ISS 19

FILE LAST UPDATED: 27 Apr 2006 (20060427/ED)

10671216

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=> s l3

L4 7 L3

=> d abs fbib fhitr 1-7

L4 ANSWER 1 OF 7 CAPLUS COPYRIGHT 2006 ACS on STN

AB A method of treating a condition comprises administering to a mammal in need of such treatment an effective amount of at least one bi- or tricyclic compound (Markush included), or a pharmaceutically acceptable isomer, salt, solvate or co-crystal form thereof. The condition is a cardiovascular or circulatory disease or condition, an inflammatory disease or condition, a respiratory tract or disease or condition, cancer, acute renal failure, astrogliosis, a fibrotic disorder of the liver, kidney, lung or intestinal tract, Alzheimer's disease, diabetes, diabetic neuropathy, rheumatoid arthritis, neurodegenerative disease, neurotoxic disease, systemic lupus erythematosus, multiple sclerosis, osteoporosis, glaucoma, macular degeneration, psoriasis, radiation fibrosis, endothelial dysfunction, a wound or a spinal cord injury, or a symptom or result thereof. Combination therapy with other therapeutically effective agents is also disclosed.

AN 2004:802569 CAPLUS

DN 141:289089

TI Methods of therapeutic use of thrombin receptor antagonists

IN Chackalamannil, Samuel; Xia, Yan; Veltri, Enrico P.; Chelliah, Mariappan V.; Wu, Wenxue; Graziano, Michael P.; Kosoglou, Teddy; Chintala, Madhu

PA USA

SO U.S. Pat. Appl. Publ., 40 pp., Cont.-in-part of U.S. Ser. No. 412,982.  
CODEN: USXXCO

DT Patent

LA English

FAN.CNT 5

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	US 2003216437	A1	20031120	US 2003-412982	20030414
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US 2003-705282

A 20031110

## PATENT FAMILY INFORMATION:

FAN 2001:923791

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JP 2004503551	T2	20040205	JP 2002-510472	20010613
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AU 2003221932	A1	20031103	WO 2003-US11510	W	20030414
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EP 1495018	A1	20050112	EP 2003-718393		20030414
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FAN 2004:633287					
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WO	2005070923	A1	20050804	WO 2005-US447	20050107
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US 2004-755066 A 20040109

OS MARPAT 141:289089

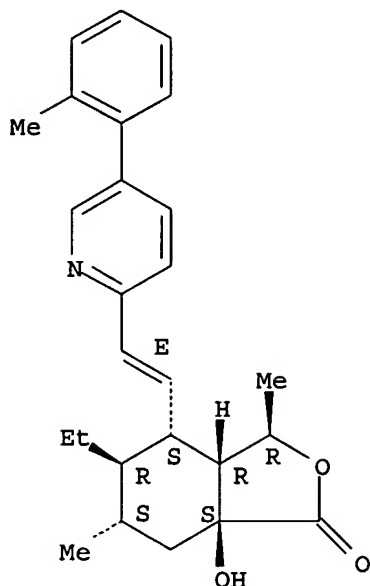
IT 380893-83-4

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL  
(Biological study); USES (Uses)  
(therapeutic use of thrombin receptor antagonists)

RN 380893-83-4 CAPLUS

CN 1(3H)-Isobenzofuranone, 5-ethylhexahydro-7a-hydroxy-3,6-dimethyl-4-[(1E)-2-[5-(2-methylphenyl)-2-pyridinyl]ethenyl]-, (3R,3aR,4S,5R,6S,7aS) - (9CI)  
(CA INDEX NAME)

Absolute stereochemistry.  
Double bond geometry as shown.



L4 ANSWER 2 OF 7 CAPLUS COPYRIGHT 2006 ACS on STN  
GI

\* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT \*

AB Eleven title compds. are claimed. Thus, phosphonate (I) in THF at 0° was treated with BuLi and then with Ti(OiPr)<sub>4</sub> followed by stirring at room temperature and addition of aldehyde (II) followed by stirring for 1.5 h to give 62% title compound (III). Title compds. inhibited thrombin receptor with IC<sub>50</sub> = 1-1000 nM.

AN 2004:633287 CAPLUS

DN 141:174354

TI Preparation of nor-seco himbacine derivatives as thrombin receptor antagonists

IN Chackalamannil, Samuel; Chelliah, Mariappan V.; Xia, Yan

PA USA

SO U.S. Pat. Appl. Publ., 52 pp., Cont.-in-part of U.S. Pat. Appl. 2004 6,105.

CODEN: USXXCO

DT Patent

LA English

FAN.CNT 5

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10671216

US 2004006105	A1	20040108	US 2000-211724P	P	20000615
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## PATENT FAMILY INFORMATION:

FAN 2001:923791

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FAN	2004:739978					
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OS MARPAT 141:174354					
IT 735287-83-9P					
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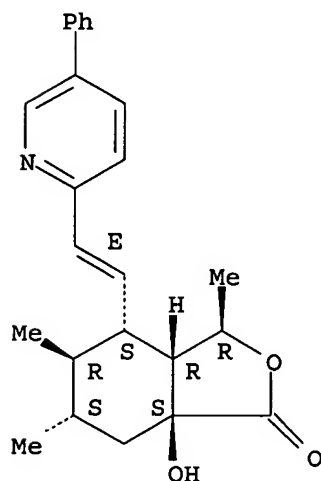
(claimed compound; preparation of nor-seco himbacine derivs. as thrombin receptor antagonists)

RN 735287-83-9 CAPLUS

CN 1(3H)-Isobenzofuranone, hexahydro-7a-hydroxy-3,5,6-trimethyl-4-[(1E)-2-(5-phenyl-2-pyridinyl)ethenyl]-, (3R,3aR,4S,5R,6S,7aS)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

Double bond geometry as shown.



L4 ANSWER 3 OF 7 CAPLUS COPYRIGHT 2006 ACS on STN  
GI

\* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT \*

AB Heterocyclic-substituted tricyclics I [B = alkylene, trans-alkenylene, alkynylene; Het = mono- or bicyclic heteroaryl; X = OH, (un)substituted NH<sub>2</sub>; XR<sub>10</sub> = O, (un)substituted NH; X<sub>1</sub> = (CH<sub>2</sub>)<sub>0-2</sub>; Y = O, S, H<sub>2</sub>, H, OH, H, alkoxy, H, SH; R<sub>1</sub>, R<sub>2</sub> = H, cycloalkyl, (un)substituted alkyl, alkenyl; R<sub>3</sub> = H, OH, alkoxy, halogen, (un)substituted alkyl, S(O)H, SO<sub>2</sub>H, CO<sub>2</sub>H, CONH<sub>2</sub>, alkenyl, cycloalkyl; R<sub>4</sub>, R<sub>7</sub>, R<sub>8</sub> = R<sub>1</sub>, OR<sub>1</sub>; R<sub>5</sub>R<sub>6</sub> = (un)substituted alkylene; R<sub>9</sub> = H, OH, alkoxy; R<sub>10</sub> = H, alkyl, (un)substituted NH<sub>2</sub>, OH] were prepared for use as thrombin receptor antagonists (no data). Thus, the naphthofuranone II [R<sub>11</sub>R<sub>12</sub> = O] was reductively aminated and treated with ClCO<sub>2</sub>Et to give II [R<sub>11</sub> = H, R<sub>12</sub> = NHCO<sub>2</sub>Et].

AN 2003:855925 CAPLUS

DN 139:350626

TI Preparation of naphthofuran thrombin receptor antagonists

IN Chackalamannil, Samuel; Clasby, Martin C.; Greenlee, William J.; Wang, Yuguang; Xia, Yan; Veltri, Enrico P.; Chelliah, Mariappan

PA Schering Corporation, USA

SO PCT Int. Appl., 64 pp.

CODEN: PIXXD2

DT Patent

LA English

10671216

## FAN.CNT 5

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PI	WO 2003089428	A1	20031030	WO 2003-US11510	20030414
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## PATENT FAMILY INFORMATION:

FAN 2001:923791

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NE, SN, TD, TG

US 2003-705282

A 20031110

OS MARPAT 139:350626

IT 618386-43-9P

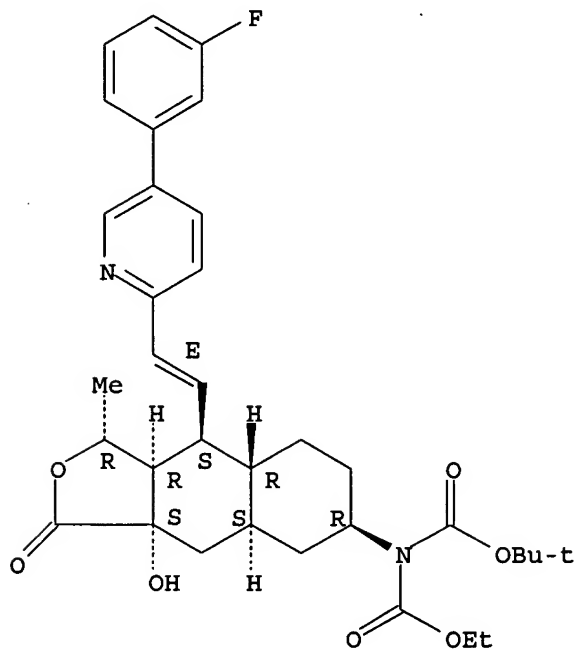
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(Reactant or reagent)  
(preparation of naphthofuran thrombin receptor antagonists)

RN 618386-43-9 CAPLUS

CN Imidodicarbonic acid, [(1R,3aS,4aS,6R,8aR,9S,9aR)-9-[(1E)-2-[5-(3-  
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INDEX NAME)

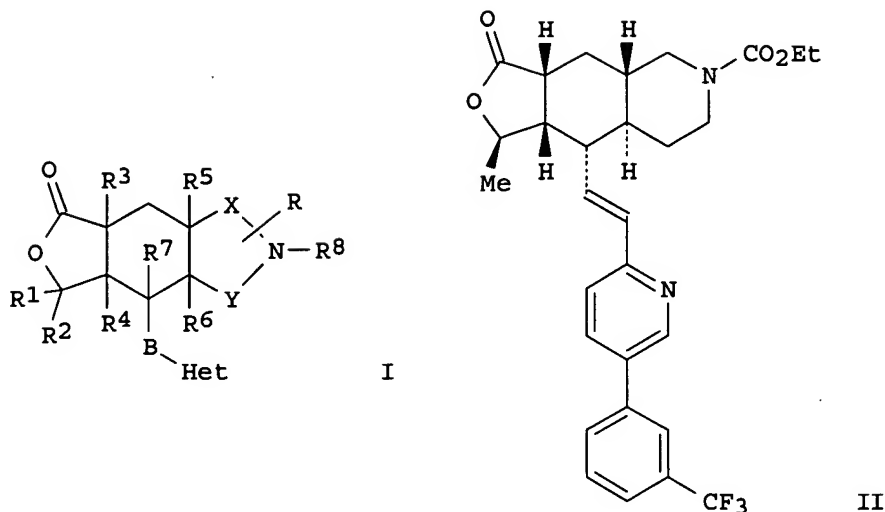
Absolute stereochemistry.

Double bond geometry as shown.



RE.CNT 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD  
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 4 OF 7 CAPLUS COPYRIGHT 2006 ACS on STN  
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AB Heterocyclic-substituted tricyclics of formula I [R = H, alkyl, halo, OH, amino, aryl, etc.; R<sup>1</sup>-R<sup>7</sup> = H, OH, alkyl, cycloalkyl, etc.; R<sup>8</sup> = acyl, carboxy, amino, etc.; X = (CH<sub>2</sub>)<sub>n</sub>; Y = (CH<sub>2</sub>)<sub>m</sub>; n, m = 0-3; B = alkyl, (substituted) alkenyl; Het = (substituted) mono-, bi- or tricyclic heteroarom. group] are prepared for treating diseases associated with thrombosis, atherosclerosis, restenosis, hypertension, angina pectoris, arrhythmia, heart failure, and cancer. Pharmaceutical compns. containing I are described. Thus, II was prepared in several steps. The prepared compds. were found to have IC<sub>50</sub> values from 1 to 2000 nM in in vivo antitumor tests against human breast carcinoma in nude mice.

AN 2003:319904 CAPLUS

DN 138:321428

TI Preparation of himbacine analogues as thrombin receptor antagonists

IN Chackalamannil, Samuel; Chelliah, Mariappan V.; Clasby, Martin C.; Xia, Yan

PA Schering Corporation, USA

SO PCT Int. Appl., 59 pp.

CODEN: PIXXD2

DT Patent

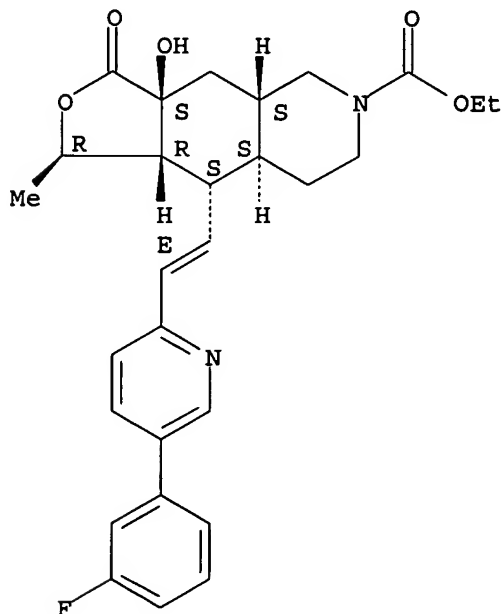
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EP 1436298	A1	20040714	EP 2002-801732	20021016
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			US 2001-330359P	P 20011018
			WO 2002-US32936	W 20021016
CN 1571789	A	20050126	CN 2002-820666	20021016
			US 2001-330359P	P 20011018
BR 2002013967	A	20050830	BR 2002-13967	20021016
			US 2001-330359P	P 20011018
			WO 2002-US32936	W 20021016
JP 2005529841	T2	20051006	JP 2003-536240	20021016
			US 2001-330359P	P 20011018
			WO 2002-US32936	W 20021016
ZA 2004002849	A	20050114	ZA 2004-2849	20040415
			US 2001-330359P	P 20011018
NO 2004002021	A	20040514	NO 2004-2021	20040514
			US 2001-330359P	P 20011018
			WO 2002-US32936	W 20021016
OS	MARPAT 138:321428			
IT	514202-92-7P			
	RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)			
	(preparation of himbacine analogs as thrombin receptor antagonists)			
RN	514202-92-7 CAPLUS			
CN	Furo[3,4-g]isoquinoline-6(3H)-carboxylic acid, 9-[(1E)-2-[5-(3-fluorophenyl)-2-pyridinyl]ethenyl]decahydro-3a-hydroxy-1-methyl-3-oxo-, ethyl ester, (1R,3aS,4aS,8aS,9S,9aR) - (9CI) (CA INDEX NAME)			

Absolute stereochemistry.  
Double bond geometry as shown.





RE.CNT 5 THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD  
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 5 OF 7 CAPLUS COPYRIGHT 2006 ACS on STN  
GI

\* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT \*

AB Title compds. I [Z = alkyl, n = 0 - 2; R1-2 = H, alk(en)yl, (di/tri)fluoroalkyl, cycloalkyl, (hetero)aryl, etc.; R3 = H, OH, alkoxy, amino, sulfinyl, sulfonyl, carboxy, etc.; Het = mono-, bi- or tricyclic heteroarom. group of 5 - 14 atoms; R8,10-11 = R1, OR1 provided that when the optional double bond is present, R10 is absent; R9 = H, OH, alkoxy, halo(alkyl); B = alkyl(oxy/thio/amino), carboxamide, etc.; X = O, N- when the double dotted line is a single bond or X = H, OH, NH- when the bond is absent; Y = O, S, (H,H), (H,OH), etc.; R15 is absent when the double dotted line represents a single bond; R15 is H, alkyl, amino or alkoxy when the bond is absent; R22-23 = H, alk(en/yn)yl, heterocycloalkyl, aryl, etc.] were prepared Over 100 synthetic examples were disclosed. E.g., II was prepared in 4 steps and reduced to the triene (THF, Lindlar's catalyst, quinoline, 1 atm H2) and subsequently heated to give a homochiral Diels-Alder adduct (m-xylene, 185°C, 10 h). Debenzylation, olefin reduction (i. EtOAc, 10% Pd/C, H2, 5 h; ii. MeOH, PtO2, 50 psi H2, 2 days) afforded the corresponding carboxylic acid (35% yield from II) and was converted to aldehyde III (CH2Cl2, ClCOCOC1, DMF; Pd(PPh3)4, (n-Bu)3SnH, 0°C, 3 h) in 48% yield. III was coupled to substituted arylmethyl diethylphosphonates (THF, n-BuLi, Ti(OPr-i)4, 0°C - room temperature) to afford example compds.; e.g. IV. Certain example compds. of the invention had IC50 = 1 - 1000 nM for the thrombin receptor. I are useful for the treatment of atherosclerosis, restenosis, hypertension, arrhythmia, etc.

AN 2001:923791 CAPLUS

DN 136:37526

TI Synthesis of  $\gamma$ -lactone-alkyl-pyridines as thrombin receptor antagonists

IN Chackalamannil, Samuel; Chelliah, Mariappan; Xia, Yan

PA Schering Corporation, USA

SO PCT Int. Appl., 69 pp.

CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 5

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 2001096330	A2	20011220	WO 2001-US19025	20010613
	WO 2001096330	A3	20020613		
	W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, HR, HU, ID, IL, IN, IS, JP, KG, KR, KZ, LC, LK, LR, LT, LU, LV, MA, MD, MG, MK, MN, MX, MZ, NO, NZ, PL, PT, RO, RU, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UZ, VN, YU, ZA			
	RW:	GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG			
				US 2000-211724P	P 20000615
	CA 2410177	AA	20011220	CA 2001-2410177	20010613
				US 2000-211724P	P 20000615

AU 2001066900	A5	20011224	WO 2001-US19025	W	20010613
			AU 2001-66900		20010613
			US 2000-211724P	P	20000615
			WO 2001-US19025	W	20010613
EP 1294714	A2	20030326	EP 2001-944492		20010613
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR					
			US 2000-211724P	P	20000615
			WO 2001-US19025	W	20010613
BR 2001011991	A	20030401	BR 2001-11991		20010613
			US 2000-211724P	P	20000615
			WO 2001-US19025	W	20010613
JP 2004503551	T2	20040205	JP 2002-510472		20010613
			US 2000-211724P	P	20000615
			WO 2001-US19025	W	20010613
NZ 523075	A	20040528	NZ 2001-523075		20010613
			US 2000-211724P	P	20000615
			WO 2001-US19025	W	20010613
NO 2002005965	A	20030214	NO 2002-5965		20021212
			US 2000-211724P	P	20000615
			WO 2001-US19025	W	20010613
ZA 2002010099	A	20040312	ZA 2002-10099		20021212
			US 2000-211724P	P	20000615

## PATENT FAMILY INFORMATION:

FAN 2003:855925

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI WO 2003089428	A1	20031030	WO 2003-US11510	20030414
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, HR, HU, ID, IL, IN, IS, JP, KG, KR, KZ, LC, LK, LR, LT, LU, LV, MA, MD, MG, MK, MN, MX, MZ, NI, NO, NZ, PH, PL, PT, RO, RU, SC, SE, SG, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UZ, VC, VN, YU, ZA, ZM				
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
			US 2002-373072P	P 20020416
CA 2482858	AA	20031030	CA 2003-2482858	20030414
			US 2002-373072P	P 20020416
			WO 2003-US11510	W 20030414
AU 2003221932	A1	20031103	AU 2003-221932	20030414
			US 2002-373072P	P 20020416
			WO 2003-US11510	W 20030414
EP 1495018	A1	20050112	EP 2003-718393	20030414
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK				
			US 2002-373072P	P 20020416
			WO 2003-US11510	W 20030414
BR 2003009309	A	20050215	BR 2003-9309	20030414
			US 2002-373072P	P 20020416
			WO 2003-US11510	W 20030414
CN 1659162	A	20050824	CN 2003-813112	20030414
			US 2002-373072P	P 20020416
JP 2005528406	T2	20050922	JP 2003-586149	20030414
			US 2002-373072P	P 20020416
			WO 2003-US11510	W 20030414
NO 2004004963	A	20041115	NO 2004-4963	20041115
			US 2002-373072P	P 20020416

FAN 2004:633287				WO 2003-US11510	W 20030414
PATENT NO.		KIND	DATE	APPLICATION NO.	DATE
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PI	US 2004152736	A1	20040805	US 2003-671216	20030925
				US 2000-211724P	P 20000615
				US 2001-880222	A3 20010613
				US 2003-457256	A2 20030609
	US 2002026050	A1	20020228	US 2001-880222	20010613
	US 6645987	B2	20031111		
				US 2000-211724P	P 20000615
	US 2004006105	A1	20040108	US 2003-457256	20030609
	US 6894065 ✓	B2	20050517		
				US 2000-211724P	P 20000615
				US 2001-880222	A3 20010613
	US 2004176418	A1	20040909	US 2004-755066	20040109
				US 2000-211724P	P 20000615
				US 2001-880222	A2 20010613
				US 2002-373072P	P 20020416
				US 2003-412982	A2 20030414
				US 2003-671216	A2 20030925
				US 2003-705282	A2 20031110
	WO 2005030712	A2	20050407	WO 2004-US31495	20040923
	W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH,				
	CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD,				
	GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC,				
	LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI,				
	NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY,				
	TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW,				
	RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM,				
	AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK,				
	EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE,				
	SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE,				
	SN, TD, TG				
				US 2003-671216	A 20030925
FAN	2004:739978				
	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
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PI	US 2004176418	A1	20040909	US 2004-755066	20040109
				US 2000-211724P	P 20000615
				US 2001-880222	A2 20010613
				US 2002-373072P	P 20020416
				US 2003-412982	A2 20030414
				US 2003-671216	A2 20030925
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	US 2002026050	A1	20020228	US 2001-880222	20010613
	US 6645987 ✓	B2	20031111		
				US 2000-211724P	P 20000615
	US 2003216437	A1	20031120	US 2003-412982	20030414
				US 2002-373072P	P 20020416
	US 2004152736	A1	20040805	US 2003-671216	20030925
				US 2000-211724P	P 20000615
				US 2001-880222	A3 20010613
				US 2003-457256	A2 20030609
	US 2004192753	A1	20040930	US 2003-705282	20031110
				US 2000-211724P	P 20000615
				US 2001-880222	A2 20010613
				US 2002-373072P	P 20020416
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WO 2005070923 A1 20050804 WO 2005-US447 20050107  
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH,  
CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD,  
GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC,  
LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI,  
NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY,  
TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW  
RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM,  
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EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, MC, NL, PL, PT,  
RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML,  
MR, NE, SN, TD, TG

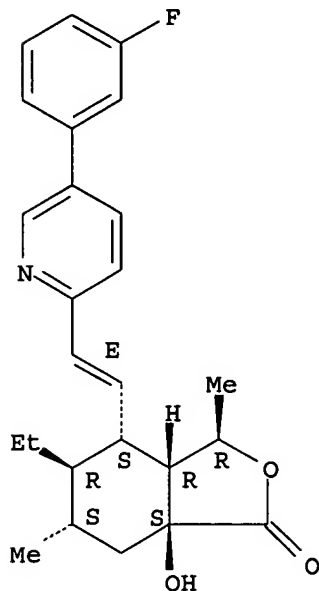
US 2004-755066 A 20040109

FAN 2004:802569  
PATENT NO. KIND DATE APPLICATION NO. DATE  
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PI US 2004192753 A1 20040930 US 2003-705282 20031110  
US 2000-211724P P 20000615  
US 2001-880222 A2 20010613  
US 2002-373072P P 20020416  
US 2003-412982 A2 20030414  
US 2001-880222 20010613  
US 2002026050 A1 20020228  
US 6645987 B2 20031111  
US 2000-211724P P 20000615  
US 2003-412982 20030414  
US 2002-373072P P 20020416  
US 2004176418 A1 20040909 US 2004-755066 20040109  
US 2000-211724P P 20000615  
US 2001-880222 A2 20010613  
US 2002-373072P P 20020416  
US 2003-412982 A2 20030414  
US 2003-671216 A2 20030925  
US 2003-705282 A2 20031110  
WO 2005046688 A2 20050526 WO 2004-US37519 20041109  
WO 2005046688 A3 20050929  
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH,  
CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD,  
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NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY,  
TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW  
RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM,  
AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK,  
EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LU, MC, NL, PL, PT, RO,  
SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR,  
NE, SN, TD, TG

US 2003-705282 A 20031110

OS MARPAT 136:37526  
IT 380893-82-3P  
RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU  
(Therapeutic use); BIOL (Biological study); PREP (Preparation); USES  
(Uses)  
(drug; synthesis of  $\gamma$ -lactone-alkyl-pyridines as thrombin  
receptor antagonists)  
RN 380893-82-3 CAPLUS  
CN 1(3H)-Isobenzofuranone, 5-ethyl-4-[(1E)-2-[5-(3-fluorophenyl)-2-  
pyridinyl]ethenyl]hexahydro-7a-hydroxy-3,6-dimethyl-,  
(3R,3aR,4S,5R,6S,7aS)-(9CI) (CA INDEX NAME)

Absolute stereochemistry.  
Double bond geometry as shown.



L4 ANSWER 6 OF 7 CAPLUS COPYRIGHT 2006 ACS on STN  
GI

\* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT \*

AB Heterocyclic-substituted tricyclics of the formula (I) [single dotted line represents an optional double bond; double dotted line represents an optional single bond; n = 0-2; Q = (un)substituted cycloalkyl, heterocycloalkyl, aryl or heteroaryl; Het = (un)substituted mono-, bi- or tricyclic heteroarom. group; B = -(CH<sub>2</sub>)<sub>n3</sub>-, wherein n<sub>3</sub> is 0-5, -CH<sub>2</sub>-O-, -CH<sub>2</sub>S-, -CH<sub>2</sub>-NR<sub>6</sub>-, -C(O)NR<sub>6</sub>-, -NR<sub>6</sub>C(O)-, etc.; X = -O- or -NR<sub>6</sub>- when the double dotted line represents a single bond, or X is -OH or -NHR<sub>20</sub> when the bond is absent; Y = =O, =S, (H, H), (H, OH) or (H, alkoxy) when the double dotted line represents a single bond, or when the bond is absent, Y = O, (H, H), (H, OH), (H, SH) or (H, C<sub>1</sub>-C<sub>6</sub> alkoxy); R<sub>15</sub> is absent when the double dotted line represents a single bond and is H, -NR<sub>18</sub>R<sub>19</sub>, or -OR<sub>17</sub> when the bond is absent; or Y = -O-(CH<sub>2</sub>)<sub>m</sub>-O- or -S-(CH<sub>2</sub>)<sub>m</sub>-S-, m = 1-2; and R<sub>15</sub>, R<sub>17</sub>, R<sub>18</sub>, R<sub>19</sub> = H or alkyl, aryl etc.] or a pharmaceutically acceptable salt were synthesized. Thus (II) was prepared starting from (R)-3-butyn-2-ol and via condensation of fragment (III) and [5-[3-(trifluoromethyl)phenyl]-2-pyridinyl]methyl-phosphonic acid di-Et ester. II shows an IC<sub>50</sub> of 0.11 nM in in vitro thrombin receptor antagonist assay. Pharmaceutical compns. containing I as well as method of treating diseases associated with thrombosis, atherosclerosis, restenosis, hypertension, angina pectoris, arrhythmia, heart failure, and cancer are described.

AN 2000:323253 CAPLUS  
DN 132:334655

TI preparation of himbacine analogs as thrombin receptor antagonists  
 IN Chackalamannil, Samuel; Asberom, Theodros; Xia, Yan; Doller, Dario;  
 Clasby, Martin C.; Czarniecki, Michael F.  
 PA Schering Corp., USA  
 SO U.S., 161 pp.  
 CODEN: USXXAM  
 DT Patent  
 LA English  
 FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	US 6063847	A	20000516	US 1998-197442	19981123
				US 1997-66518P	P 19971125
	US 6326380	B1	20011204	US 2000-545720	20000407
				US 1997-66518P	P 19971125
				US 1998-197442	A3 19981123

OS MARPAT 132:334655

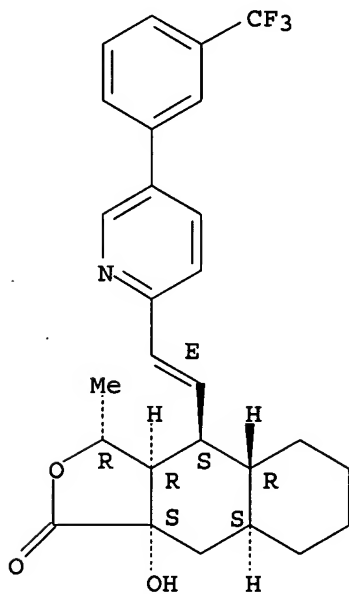
IT 226914-44-9P

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)  
 (preparation of himbacine analog as thrombin receptor antagonists)

RN 226914-44-9 CAPLUS

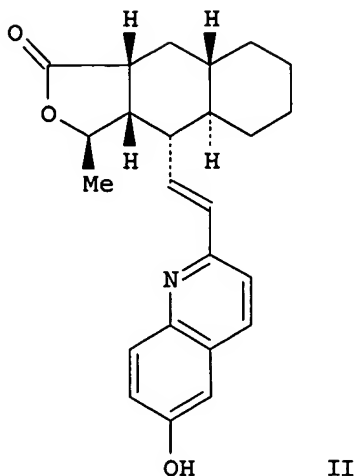
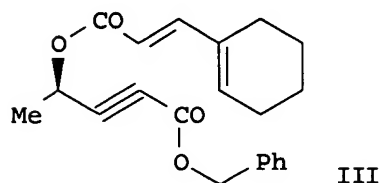
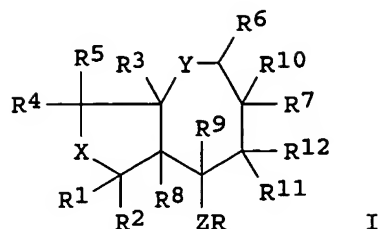
CN Naphtho[2,3-c]furan-1(3H)-one, decahydro-9a-hydroxy-3-methyl-4-[(1E)-2-[5-[3-(trifluoromethyl)phenyl]-2-pyridinyl]ethenyl]-, (3R,3aR,4S,4aR,8aS,9aS)-(9CI) (CA INDEX NAME)

Absolute stereochemistry.  
 Double bond geometry as shown.



RE.CNT 11 THERE ARE 11 CITED REFERENCES AVAILABLE FOR THIS RECORD  
 ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 7 OF 7 CAPLUS COPYRIGHT 2006 ACS on STN  
 GI



AB Himbacine analogs I [R = heteroaryl, such as pyridinyl, quinolinyl, isoquinolinyl, etc; R1, R2, R8, R10, R11 = H, alkyl, fluoroalkyl, cycloalkyl, alkenyl, aryl, heteroaryl, etc.; R3 = H, OH, alkoxy, alkylsulfinyl, alkylsulfonyl, alkyl, carboxyl, carbamido, aryl, etc.; R4, R5 = H, OH, alkyloxy, alkyl, amino, etc.; R4R5 = O, S; R6 = H; R6R10 = bond; R7R12 = fused alicyclic, fused aryl, fused heteroaryl, etc.; R9 = H, OH, alkoxy, halogen, haloalkyl; X = O, NR13; R13 = H, alkyl, Ph, etc.; Y = (CH2)n, n = 0 - 2; Z = connecting group, such as CH:CH, CH2CH2, CH2O, CH2S, CH2NH, CONH, etc.] were prepared for use as thrombin receptor antagonists for the treatment of diseases associated with thrombosis, atherosclerosis, restenosis, hypertension, angina pectoris, arrhythmia, heart failure, and cancer. Thus, lactone II was prepared starting from (R)-3-butyn-2-ol, trans-3-(1-cyclohexenyl)acrylic acid, and 6-hydroxyquinoline via the formation and intramol. cycloaddn. of diester III. The prepared compds. were tested for thrombin receptor binding, platelet aggregation and antitumor activity.

AN 1999:355771 CAPLUS

DN 131:32085

TI Preparation of himbacine analogs for use as thrombin receptor antagonists

IN Chackalamannil, Samuel; Asberom, Theodoros; Xia, Yan; Doller, Dario;

Clasby, Martin C.; Czarniecki, Michael F.

PA Schering Corporation, USA

SO PCT Int. Appl., 134 pp.

CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 9926943	A1	19990603	WO 1998-US24523	19981123
	W: AL, AM, AU, AZ, BA, BB, BG, BR, BY, CA, CN, CZ, EE, GD, GE, HR, HU, ID, IL, IS, JP, KG, KR, KZ, LC, LK, LR, LT, LV, MD, MG, MK, MN, MX, NO, NZ, PL, RO, RU, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UZ, VN, YU, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
	RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI,				

CM, GA, GN, GW, ML, MR, NE, SN, TD, TG					
			US 1997-977979	A	19971125
CA 2309352	AA	19990603	CA 1998-2309352		19981123
CA 2309352	C	20050125			
			US 1997-977979	A	19971125
			WO 1998-US24523	W	19981123
AU 9914158	A1	19990615	AU 1999-14158		19981123
AU 747204	B2	20020509			
			US 1997-977979	A	19971125
			WO 1998-US24523	W	19981123
ZA 9810685	A	19991223	ZA 1998-10685		19981123
			US 1997-977979	A	19971125
EP 1036072	A1	20000920	EP 1998-958039		19981123
EP 1036072	B1	20040506			
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, PT, IE, LT, LV, FI, RO					
			US 1997-977979	A	19971125
			WO 1998-US24523	W	19981123
TR 200001480	T2	20000921	TR 2000-200001480		19981123
			US 1997-977979	A	19971125
BR 9812793	A	20001017	BR 1998-12793		19981123
			US 1997-977979	A	19971125
			WO 1998-US24523	W	19981123
JP 2001524479	T2	20011204	JP 2000-522101		19981123
JP 3449620	B2	20030922			
			US 1997-977979	A	19971125
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JP 2003128670	A2	20030508	JP 2002-315015		19981123
			US 1997-977979	A	19971125
			JP 2000-522101	A3	19981123
RU 2204557	C2	20030520	RU 2000-116548		19981123
			US 1997-977979	A	19971125
			WO 1998-US24523	W	19981123
IL 135797	A1	20030917	IL 1998-135797		19981123
			US 1997-977979	A	19971125
			WO 1998-US24523	W	19981123
AT 266025	E	20040515	AT 1998-958039		19981123
			US 1997-977979	A	19971125
			WO 1998-US24523	W	19981123
PT 1036072	T	20040831	PT 1998-958039		19981123
			US 1997-977979	A	19971125
ES 2219919	T3	20041201	ES 1998-958039		19981123
			US 1997-977979	A	19971125
NO 2000002659	A	20000724	NO 2000-2659		20000524
			US 1997-977979	A	19971125
			WO 1998-US24523	W	19981123
HK 1031726	A1	20040930	HK 2001-101899		20010316
			US 1997-977979	A	19971125
			WO 1998-US24523	W	19981123
OS	MARPAT 131:32085				
IT	226914-44-9P				
	RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)				
	(preparation of himbacine analogs for use as thrombin receptor antagonists)				
RN	226914-44-9 CAPLUS				
CN	Naphtho[2,3-c]furan-1(3H)-one, decahydro-9a-hydroxy-3-methyl-4-[(1E)-2-[5-[3-(trifluoromethyl)phenyl]-2-pyridinyl]ethenyl]-, (3R,3aR,4S,4aR,8aS,9aS)-(9CI) (CA INDEX NAME)				



Absolute stereochemistry.  
Double bond geometry as shown.

